FIGURED.



1647

180% JEWAH 1801 2900

RAW SEQUENCE LISTING DATE: 03/28/2002 PATENT APPLICATION: US/09/903,180B TIME: 10:51:04

| | <110> | | | | | | is, I | Edwa | rd M | | | | | | | ļ., | ECEIVED |
|----------|-------|---|-----------|----------|-----------|----------|-------|------------|--------|------------------|-------|------|-------|---------|------------------|-------|---------|
| 2 | -100- | | | | | | | | | | | | | | | | |
| | <120> | > TITLE OF INVENTION: Endoderm, Cardiac and Neural Inducing | | | | | | | | | | | | ·, 18.5 | | | |
| 4 | -120 | Factors | | | | | | | | | | | | | ÷ - 2+ | | |
| | | FILE REFERENCE: 510015-256 | | | | | | | | | | | | | CE: ER: 800 2900 | | |
| | | CURRENT APPLICATION NUMBER: US/09/903,180B CURRENT FILING DATE: 2001-07-11 | | | | | | | | | | | | | - V | | |
| | | PRIOR APPLICATION NUMBER: US 60/020,150 | | | | | | | | | | | | | | | |
| | | | | | | | | | | J Z U , . | 130 | | | | _ | | |
| | <151> | | | | | | | J 6 - Z (| J | | | | | | | `A. | |
| | | NUMBER OF SEQ ID NOS: 10 SOFTWARE: FastSEQ for Windows Version 3.0 | | | | | | | | | | | | | | Tr | |
| | | " <i>F.</i> 1 | | | | | | | | | | | | | | | |
| | | SEQ ID NO: 1 | | | | | | | | | | | | | | | |
| | | LENGTH: 270 | | | | | | | | | | | | | | | |
| | | TIPE. PRI | | | | | | | | | | | | | | | |
| | | > ORGANISM: Xenopus > SEQUENCE: 1 | | | | | | | | | | | | | | | |
| | <400> | | | | Asn | 37 n] | T 0 | 7 | т1. | Crra | т1о | Tlo | Wa 2 | Cvrc | T OU | Wa l | λan |
| 18 | | | Leu | Leu | ASII | vai 5 | Leu | Arg | TIE | Cys | 10 | ше | vai | Cys | ьеи | 15 | ASII |
| 19 | | 1 | ~1 | 7. T. n. | C1 | _ | ni a | Cox | C1., | C1,, | | Clu | Λrα | Thr | Tvc | | Tur |
| 20 | | ASP | GIY | Ата | Gly | гуѕ | птѕ | ser | GIU | 25 | Alg | Giu | Arg | 1 111 | 30 | 1111 | 171 |
| 21 | | C = == | т о | n an | 20 Ser | N mar | C1., | Пттх | Dho | | Tvc | Clu | λνα | C1 17 | | λνα | Ara |
| 22 | | ser | Leu | 35 | ser | AIG | СТУ | тут | 40 | AIG | гуз | Gru | AIG | 45 | AIG | Alg | hig |
| 23 | | C 0 m | T *** | | Leu | T 011 | W > 1 | N a n | | Lvc | Clu | Lou | Nan | | Dro | ије | Tlα |
| 24 | | Ser | БуS 50 | 116 | ьeu | Leu | vai | 55 | 1111 | гуз | Gry | Leu | 60 | GIU | FIO | 1113 | 110 |
| 25 | | C1 | | c1 | Asp | Dho | C1 | | 17 - 1 | λla | Clu | LOU | | λen | Car | Thr | λra |
| 26 | | 65 | птъ | GTA | ASP | PHE | 70 | Leu | Val | нта | Giu | 75 | FIIC | изр | DCI | 1111 | 80 |
| 27 | | | II d | mb x | Asn | λrα | | Clu | Dro | λcn | Mot | | Luc | Va 1 | Tave | T.eu | |
| 28 29 | | 1111 | HIS | 1111 | ASII | 85 | БУЗ | Giu | PIO | АЗР | 90 | NSII | цуз | vai | цуз | 95 | THE |
| | | Cor | πh∽ | Val | Ala | | C117 | λan | Lvc | Sor | | λra | Δra | Luc | Δla | | Δen |
| 30 | | ser | 1 111 | val | 100 | птъ | GIY | ASII | гуз | 105 | AIG | Arg | AIG | цуз | 110 | 1 Y L | ASII |
| 31 32 | | C1 v | cor | λrα | Arg | Nen | Tla | Dha | Sor | | Δνα | Ser | Dhe | Δsn | | Ara | Agn |
| 33 | | GIY | ser | 115 | Alg | ASII | 116 | FIIC | 120 | Arg | Arg | JCI | 1 110 | 125 | БуЗ | 1119 | 11511 |
| 34 | | Thr | Clu | | Thr | Glu | Twe | Dro | | Δla | T.375 | Met | Dhe | | Asn | Δsn | Phe |
| 35 | | 1111 | 130 | Val | 1111 | Giu | цуз | 135 | GIY | AIU | цуз | ricc | 140 | 111 | 11511 | 11511 | 1110 |
| 36 | | Lou | | Two | Met | λen | C1v | | Dro | Gln | Δen | Thr | | His | Glv | Ser | Lvs |
| 37 | | 145 | VUI | цуз | Mec | ASII | 150 | niu | 110 | 0111 | 11011 | 155 | DCI | | 0 - 1 | 001 | 160 |
| 38 | | | C1n | Clu | Ile | Mot | | Glu | Δla | Cvs | T.vc | | Len | Pro | Phe | Thr | |
| 39 | | AId | GIII | GIU | 116 | 165 | цуз | GIU | лıu | Cys | 170 | 1111 | Leu | 110 | 1 110 | 175 | J |
| 40 | | Aen | Tle | Va 1 | His | | Aen | Cvc | Aan | Ara | | Val | Tle | Gln | Asn | | Leu |
| 41 | | บอเเ | 116 | v a I | 180 | GIU | บอน | Cys | rab | 185 | 1100 | · uı | 110 | 0111 | 190 | 11011 | |
| 42 | | Cve | Dhe | Glv | Lys | Cvc | Tle | Ser | Len | | Va 1 | Pro | Asn | Gln | | Asp | Ara |
| 43 | | Cys | 1 116 | 195 | Lys | CyB | 110 | 001 | 200 | | , 41 | | | 205 | o | | J |
| 44 | | Ara | Asn | | Cys | Ser | His | Cvs | | Pro | Ser | Lvs | Phe | | Leu | Asn | His |
| 77 | | Arg | asa | 1111 | CIB | 501 | **** | $\cup_I S$ | a | 0 | 501 | 2,5 | | | | | |

RAW SEQUENCE LISTING

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| 45 | 210 | | 215 | | 220 | |
|----|------------------|-------------|--------------|--------------|-----------------------|------|
| 46 | | eu Asn Cvs | | Lys Asn Val | Val Lys Val Val Met | |
| 47 | 225 | -1- | 230 | 235 | 240 | |
| 48 | | lu Glu Cys | Thr Cys Glu | Ala His Lys | Ser Asn Phe His Gln | |
| 49 | | 245 | | 250 | 255 | |
| 50 | Thr Ala G | ln Phe Asn | Met Asp Thi | Ser Thr Thr | Leu His His | |
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| 60 | | | | | aggttacttc agaaaagaaa | 180 |
| 61 | | | | | aggtcttgat gaaccccaca | 240 |
| 62 | | | | | ttccaccaga acacatacaa | 300 |
| 63 | | | | | aacagttgcc catggaaaca | 360 |
| 64 | | | | | tatttttcct cgccgttctt | 420 |
| 65 | ttgataaaa | g aaatacag | ag gttactgaa | a agcctggtgc | caagatgttc tggaacaatt | 480 |
| 66 | | | | | tggcagtaaa gcacaggaaa | 540 |
| 67 | taatgaaag | a agcttgca | aa accttgttt | t tcactcagaa | tattgtacat gaaaactgtg | 600 |
| 68 | | | | | catctctctc catgttccaa | 660 |
| 69 | | | | | gtccaaattt accctgaacc | 720 |
| 70 | acctgacgo | t gaattgta | ct ggatctaag | a atgtagtaaa | ggttgtcatg atggtagagg | 780 |
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| 73 | | - | | - | cttctatttc atataaccac | 960 |
| 74 | atggaataa | g gattgtat | ga attataatt | a acaaatggca | ttttgtgtaa catgcaagat | 1020 |
| 75 | ctctqttcc | a tcagttgc | aa gataaaagg | c aatatttgtt | tgacttttt tctacaaaat | 1080 |
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| 78 | | | | | gggttactgc ttctgggcaa | 1260 |
| 79 | aatgtgtgc | c tcataggg | gg gttagtgtg | t tgtgtactga | ataaattgta tttatttcat | 1320 |
| 80 | tgttacaaa | a aaaaaaaa | | | | 1338 |
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| 89 | Gly Leu A | la Leu Leu | Leu Leu Pro | Asn Ala Tyr | Cys Ala Ser Cys Glu | |
| 90 | | 20 | | 25 | 30 | |
| 91 | | - | | Ser Met Pro | Trp Asn Met Thr Lys | |
| 92 | | 5 | 40 | _, _, _ | 45 | |
| 93 | | sn His Leu | | Thr Gln Ala | Asn Ala Ile Leu Ala | |
| 94 | 50 | | 55 | -1 -1 -1 | 60 | |
| 95 | Ile Glu G | In Phe Glu | Gly Leu Leu | Thr Thr Glu | Cys Ser Gln Asp Leu | |

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| 97 | | Phe | Phe | Leu (| | | Met ' | Tvr | Ala | Pro | | Cvs | Thr | Ile | Asp | | |
| 98 | Dea | 1110 | | | 85 | | | -1- | | 90 | | -1- | | | 95 | | |
| 99 | Gln | His | Glu | | | Lvs | Pro (| Cvs : | Lvs | | Val | Cvs | Glu | Arq | Ala | Arq | |
| 100 | 01 | | 0 | 100 | | -1- | | -1- | 105 | | | . 4 | | 110 | | , | |
| 101 | Ala | Gly | Cvs | | Pro | Ile | Leu | Ile | | | Aro | His | Thr | Trp | Pro | Glu | |
| 102 | 1120 | . 011 | 115 | | | | | 120 | -1- | - 1 | 2 | | 125 | | | | |
| 103 | Ser | Leu | | | Glu | Glu | Leu | | Val | Tvr | Asp | Arq | Glv | Val | Cvs | Ile | |
| 104 | 001 | 130 | | • ₁ = | | | 135 | | | - 1 - | - 1 | 140 | | | - | | |
| 105 | Ser | | | Ala | Ile | Val | | Val | Glu | Gln | Gly | Thr | Asp | Ser | Met | Pro | |
| 106 | 145 | | | | | 150 | | | | | 155 | | - | | | 160 | |
| 107 | | | Ser | Met | Asp | | | Asn | Glv | Asn | Cys | Gly | Ser | Gly | Arq | Glu | |
| 108 | | | | | 165 | | | | 1 | 170 | | | | _ | 175 | | |
| 109 | His | . Cys | Lvs | Cvs | | Pro | Met | Lvs | Ala | Thr | Glr | Lys | Thr | Tyr | Leu | Lys | |
| 110 | | | -1- | 180 | -1 - | | | _1 | 185 | | | • | | 190 | | - | |
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| 112 | | | 195 | | -1- | | | 200 | | 1 | | - | 205 | | _ | | |
| 113 | Lvs | . Cys | | | Ala | Thr | Ala | | Val | Glu | Val | Lvs | | | Leu | Lys | |
| 114 | -1- | 210 | | | | - | 215 | | | | | 220 | | | | - | |
| 115 | Ser | | | Val | Asn | Ile | | Lvs | Asp | Thr | Val | Thr | Leu | Tyr | Thr | Asn | |
| 116 | 225 | | | | | 230 | | -1- | | | 235 | | | • | | 240 | |
| 117 | | | Cvs | Leu | Cvs | | | Leu | Val | Ala | | | Glu | Tvr | · Ile | lle | |
| 118 | 001 | . 011 | 010 | | 245 | | | | | 250 | | | | 1 | 255 | | |
| 119 | Met | Glv | Tvr | Glu | | Lvs | Glu | Arg | Thr | | | Leu | Leu | Val | Glu | Gly | |
| 120 | 1100 | . 011 | +1- | 260 | E | -1- | | 5 | 265 | | | | | 270 | | _ | |
| 121 | Ser | · Leu | Ala | | Lvs | Trp | Ara | Asp | | | Ala | Lvs | Lvs | Val | Lys | Arg | |
| 122 | | | 275 | | -1- | | 5 | 280 | 5 | | | 4 | 285 | | - | , | |
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| | 11> LENG | | | | | | | | | | | | | | | | |
| | 12> TYPE | | | | | | | | | | | | | | | | |
| | 13> ORG <i>A</i> | | | nopu | s fr | azzl | ed | | | | | | | | | | |
| | 00> SEQU | | | • | | | | | | | | | | | | | |
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| 139 | | | | | | | | | | | | | | | | .cgtgt | |
| 140 | ago | ctgt | gcg | gatc | ccca | tġ ť | gcaa | atct | a tg | ccat | ggaa | cat | .gacc | aag | atgo | ccaac | c 480 |
| 141 | | | | | | | | | | | | | | | | gtttg | |
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| 144 | | | | | | | | | | | | | | | | tggca | |
| 145 | | | | | | | | | | | | | | | | rtcaca | |
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| 146 | | | | | | | | | | | | | | | | | | | 0.40 |
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| 148 | | | | | | | | | | | | | | | | | | | |
| 149 | | | | | | | | | | | | | | | | | | | |
| 150 | | | _ | | | | _ | | _ | - | | | _ | | | | | | |
| 151 | | | | | | | | | | | | | | | | | | | |
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| 156 | | | | | | | | | | | | | | | | | | | |
| 157 | | | | _ | | _ | _ | _ | | - | | | | | | | | | |
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| 160 | | | | | | | | | | | | | | | | | | | |
| 161 | | | | | | | | | | | | | | | | | | | |
| 1800 163 | | | | | - | | _ | - | _ | _ | _ | | | | | | | | |
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| 170 | | | | | | | | | | | | | | | | | | | |
| 171 | | | | | | nopus | S | | | | | | | | | | | | |
| 172 | 170 | <400> | | | | | | | | | | | | | | | | | |
| 173 | 171 | | Met | Leu | Leu | Leu | Phe | Arg | Ala | Ile | Pro | Met | Leu | Leu | Leu | Gly | | Met | |
| 174 | 172 | | _ | | | | - | | | | | | | | | | | | |
| 175 | 173 | | Val | Leu | Gln | | Asp | Cys | Glu | Ile | Ala | Gln | Tyr | Tyr | Ile | | Glu | Glu | |
| 176 177 188 179 180 180 180 181 180 181 180 182 183 183 184 185 186 187 188 188 188 188 188 188 188 188 188 | | | | | | | | | | | | | | | | | _ | _ | |
| 177 | | | Glu | Pro | | Gly | Thr | Val | Ile | | Val | Leu | Ser | Gln | | Ser | Ile | Phe | |
| 178 50 55 60 179 Asn Asn Ser Asn Ser Leu Ile Gly Val Arg Glu Ser Asp Gly Gln Leu Ser Ile 80 180 65 70 70 80 80 181 Met Glu Arg Ile Asp Ser Ile 85 | | | | | | | | | | | | _ | | | | | | | |
| 179 | | | Asn | | Thr | Asp | Ile | Pro | | Thr | Asn | Phe | Arg | | Met | Lys | GIn | Phe | |
| 180 65 | | | | | | | | _ | | | | | | | | _ | _ | - 2 | |
| 181 Met Glu Arg Ile Asp St Arg Glu Gln Ile Cys Arg Gln Ser Leu His Cys 90 Ser Leu His Cys 95 182 85 90 90 95 183 Asn Leu Ala Leu Asp Val Val Ser Phe Ser Lys Gly His Phe Lys Leu 100 110 110 184 100 105 110 110 185 Leu Asn Val Lys Val Glu Val Arg Asp Ile Asn Asp His Ser Pro His 125 125 125 186 115 120 125 125 187 Phe Pro Ser Glu Ile Met His Val Glu Val Ser Glu Ser Ser Ser Val 135 140 188 130 135 140 189 Gly Thr Arg Ile Pro Leu Glu Ile Ala Ile Asp Glu Asp Val Gly Ser 150 155 160 191 Asn Ser Ile Gln Asn Phe Gln Ile Ser Asn Asn Ser His Phe Ser Ile 165 170 175 193 Asp Val Leu Thr Arg Ala Asp Gly Val Lys Tyr Ala Asp Leu Val Leu 190 190 | | | | Asn | Ser | Leu | Ile | | Val | Arg | Glu | Ser | | GLY | GIn | Leu | Ser | | |
| 182 Asn Leu Ala Leu Asp Val Val Ser Phe Ser Lys Gly His Phe Leu Lys Leu Leu Asp Val Val Ser Phe Ser Lys Leu Asp Val Leu Asp Ile Asp Ile Asp Ile Asp His Ser Pro His Ile His Ile Asp Ile Asp Ile Asp His Ser Pro His Ile I | | | | _ | | _ | | | | | | _ | | | _ | _ | | | |
| Asn Leu Ala Leu Asp Val Val Ser Phe Ser Lys Gly His Phe Lys Leu 105 | | | Met | Glu | Arg | Ile | | Arg | Glu | GIn | He | | Arg | GIn | Ser | Leu | | Cys | |
| 184 100 105 110 185 Leu Asn Val Lys Val Glu Val Arg Asp Ile Asn Asp His Ser Pro His 186 115 120 125 187 Phe Pro Ser Glu Ile Met His Val Glu Val Ser Glu Ser Ser Ser Val 188 130 135 140 189 Gly Thr Arg Ile Pro Leu Glu Ile Ala Ile Asp Glu Asp Val Gly Ser 190 145 150 155 160 191 Asn Ser Ile Gln Asn Phe Gln Ile Ser Asn Asn Ser His Phe Ser Ile 165 170 175 193 Asp Val Leu Thr Arg Ala Asp Gly Val Lys Tyr Ala Asp Leu Val Leu 190 185 190 | | | | | | _ | | | | _ | _, | | _ | a 1 | | 51 | | | |
| Leu Asn Val Lys Val Glu Val Arg Asp Ile Asn Asp His Ser Pro His 186 115 120 125 187 Phe Pro Ser Glu Ile Met His Val Glu Val Ser Glu Ser Ser Ser Val 188 130 130 135 140 189 Gly Thr Arg Ile Pro Leu Glu Ile Ala Ile Asp Glu Asp Val Gly Ser 190 145 150 155 160 191 Asn Ser Ile Gln Asn Phe Gln Ile Ser Asn Asn Ser His Phe Ser Ile 192 165 170 175 189 Asp Val Leu Thr Arg Ala Asp Gly Val Lys Tyr Ala Asp Leu Val Leu 194 | | | Asn | Leu | Ala | | Asp | Val | Val | Ser | | Ser | Lys | GLY | HIS | | Lys | Leu | |
| 186 115 120 125 187 Phe Pro Ser Glu Ile Met His Val Glu Val Ser Glu Ser Ser Ser Val 188 130 135 140 189 Gly Thr Arg Ile Pro Leu Glu Ile Ala Ile Asp Glu Asp Val Gly Ser 190 145 150 155 160 191 Asn Ser Ile Gln Asn Phe Gln Ile Ser Asn Asn Ser His Phe Ser Ile 165 170 175 193 Asp Val Leu Thr Arg Ala Asp Gly Val Lys Tyr Ala Asp Leu Val Leu 190 185 190 | | | | | _ | | | | | | | | _ | _ | • | | _ | | |
| 187 Phe Pro Ser Glu Ile Met His Val Glu Val Ser Glu Ser Ser Ser Val 188 130 135 140 189 Gly Thr Arg Ile Pro Leu Glu Ile Ala Ile Asp Glu Asp Val Gly Ser 190 145 150 155 160 191 Asn Ser Ile Gln Asn Phe Gln Ile Ser Asn Asn Ser His Phe Ser Ile 165 170 175 193 Asp Val Leu Thr Arg Ala Asp Gly Val Lys Tyr Ala Asp Leu Val Leu 190 185 190 | | | Leu | Asn | | Lys | Val | Glu | Val | | Asp | Ile | Asn | Asp | | Ser | Pro | HlS | |
| 188 | | | | | | _ | | | | | | | _ | 1 | | _ | _ | 1 | |
| 189 Gly Thr Arg Ile Pro Leu Glu Ile Ala Ile Asp Glu Asp Val Gly Ser 190 145 150 155 160 191 Asn Ser Ile Gln Asn Phe Gln Ile Ser Asn Asn Ser His Phe Ser Ile 192 165 170 175 193 Asp Val Leu Thr Arg Ala Asp Gly Val Lys Tyr Ala Asp Leu Val Leu 194 180 185 190 | | | Phe | | Ser | Glu | Ile | Met | | Val | Glu | Val | Ser | | Ser | Ser | Ser | Val | |
| 190 145 150 155 160 191 Asn Ser Ile Gln Asn Phe Gln Ile Ser Asn Asn Ser His Phe Ser Ile 192 165 170 175 193 Asp Val Leu Thr Arg Ala Asp Gly Val Lys Tyr Ala Asp Leu Val Leu 194 180 185 195 190 | | | | | | | | | | | _ | _ | | | | | | | |
| Asn Ser Ile Gln Asn Phe Gln Ile Ser Asn Asn Ser His Phe Ser Ile 192 165 170 175 193 Asp Val Leu Thr Arg Ala Asp Gly Val Lys Tyr Ala Asp Leu Val Leu 194 180 185 190 | | | _ | Thr | Arg | Ile | Pro | | Glu | Ile | Ala | Ile | | Glu | Asp | Val | Gly | | |
| 192 165 170 175 193 Asp Val Leu Thr Arg Ala Asp Gly Val Lys Tyr Ala Asp Leu Val Leu 194 180 185 190 | | | | | | | | | | | | | | _ | | | _ | | |
| 193 Asp Val Leu Thr Arg Ala Asp Gly Val Lys Tyr Ala Asp Leu Val Leu 194 180 185 190 | | | Asn | Ser | Ile | Gln | | Phe | Gln | Ile | Ser | | Asn | Ser | His | Phe | | пе | |
| 194 180 185 190 | | | | | _ | | | | _ | | ~ | | _ | | _ | _ | | _ | |
| | | | Asp | Val | Leu | | Arg | Ala | Asp | Gly | | Lys | Tyr | Ala | Asp | | | Leu | |
| 195 Met Arg Glu Leu Asp Arg Glu Ile Gln Pro Thr Tyr Ile Met Glu Leu | | | | _ | | | _ | _ | | | | _ | ~1 | _ | | | | - | |
| | 195 | | Met | Arg | Glu | Leu | Asp | Arg | Glu | Пе | GIn | Pro | Thr | Tyr | шe | мet | Glu | ьeu | |